

WHAT IS CLAIMED IS:

1           1. A method for comparing a test genome to a reference genome, said  
2 method comprising:

3               (i) providing a plurality of clones of known size that substantially cover at  
4 least a portion of said test genome;

5               (ii) obtaining sequence information from the termini of each of said  
6 plurality of clones, thereby obtaining a pair of terminal sequences;

7               (iii) identifying a pair of sequences within said reference genome that  
8 corresponds to each of said pairs of terminal sequences; and

9               (iv) determining the relationship between the members of each pair of  
10 corresponding sequences within said reference genome;

11               wherein a difference in the observed relationship between the members of  
12 any of said pairs of corresponding sequences within said reference genome and the  
13 expected relationship based upon said known size of said plurality of clones indicates the  
14 presence of a rearrangement in said test genome compared to said reference genome.

1           2. The method of claim 1, further comprising determining the  
2 sequence of said test genome over a region spanning at least one breakpoint of said  
3 rearrangement.

1           3. The method of claim 1, wherein said reference genome is a human  
2 genome.

1           4. The method of claim 1, wherein said test genome is from a tumor  
2 cell.

1           5. The method of claim 1, wherein said reference genome and said  
2 test genome are from different species.

1           6. The method of claim 1, wherein said plurality of clones covers  
2 substantially all of said test genome.

1           7. The method of claim 1, wherein the members of at least one pair of  
2 corresponding sequences within said reference genome are closer together than expected  
3 based on said known size of said plurality of clones, indicating the presence of an

4 insertion in said test genome between the pair of terminal sequences corresponding to  
5 said at least one pair of corresponding sequences.

1 8. The method of claim 1, wherein the members of at least one pair of  
2 corresponding sequences within said reference genome are further apart than expected  
3 based on said known size of said plurality of clones, indicating the presence of a deletion  
4 in said test genome between the pair of terminal sequences corresponding to said at least  
5 one pair of corresponding sequences.

1 9. The method of claim 1, wherein the members of at least one pair of  
2 corresponding sequences within said reference genome are present on different  
3 chromosomes within said reference genome, indicating the presence of a translocation in  
4 said test genome between the pair of terminal sequences corresponding to said at least one  
5 pair of corresponding sequences.

1 10. The method of claim 1, further comprising determining the  
2 frequency of each of said terminal sequences within said plurality of clones, wherein an  
3 increased or decreased relative frequency of any of said terminal sequences indicates the  
4 presence of an amplification or a deletion in said test genome that includes said terminal  
5 sequence.

1 11. The method of claim 1, wherein said plurality of clones are BAC  
2 clones.

1 12. The method of claim 1, wherein said plurality of clones are PAC  
2 clones.

1 13. The method of claim 1, wherein said plurality of clones represents  
2 a redundancy of at least about 10 fold of said test genome or said portion of said test  
3 genome.

1 14. The method of claim 13, wherein said plurality of clones represents  
2 a redundancy of at least about 20 fold of said test genome or said portion of said test  
3 genome.

1           15. The method of claim 1, wherein said terminal sequences are  
2 present on average between about every 5 kb to about every 500 kb throughout said test  
3 genome or said portion of said test genome.

1           16. The method of claim 15, wherein said terminal sequences are  
2 present on average every 50 kb or less throughout said test genome or said portion of said  
3 test genome.

1           17. The method of claim 16, wherein said terminal sequences are  
2 present on average every 10 kb or less throughout said test genome or said portion of said  
3 test genome.

1           18. The method of claim 17, wherein said terminal sequences are  
2 present on average every 5 kb or less throughout said test genome or said portion of said  
3 test genome.

1           19. The method of claim 1, wherein said reference genome is a human  
2 genome, and wherein said plurality of clones comprises at least about 100,000 clones.

1           20. The method of claim 19, wherein said plurality of clones comprises  
2 at least about 200,000 clones.

1           21. The method of claim 20, wherein said plurality of clones comprises  
2 at least about 250,000 clones.

1           22. The method of claim 1, wherein said terminal sequences are  
2 determined by automated sequencing.

1           23. The method of claim 1, wherein said pairs of terminal sequences  
2 from said test genome are compared to said pairs of corresponding sequences within said  
3 reference genome using a computer.

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